

Fig. 1

TGGTTGTCCTGGAACCTCACTCTGTAGACCAGGCTGGCCATGAACTCACAGA
GATCTACCTCCTGAGTGCTGGGATTAAAGGTTTGTGCCACCACCTCCCAACT
CTAAGGTGTTTCTTTAAGTTAGGGGCATAGTAAACATTGTTGAGATACTAGA
GGAACACTGAATGAAAATTGGACATCTCTGCTTTAGGTTTGTGCTGAGCA
GTTTGCCTCTTATCTTCACCTATGCTGAAAAGTTTGAGTTCATAATTTTGAA
CATGCATATGATAAAATATTCTGGCCGCACATTGAATAAATATATTTTAAAT
GAACTTACCTTTAAAATGTCAGTAACAACCTCTGCATGGTTTTCTTCTTACCT
CCATAGGTATGGTCTGAATATGCGTTGTTTGGCAGCTCGGGTCAACTATAA
GACTTTGATTATCATCTGTGCGCTATTCACTTTGGTCACAGTACTTTTGTGG
AATAAGTGTTCCAGCGACAAAGCAATCCAGTTTCCTCGGCACTTGAGTAGT
GGATTCAGAGTGATGGATTAGAAAAAAGATCAGCAGCATCTGAAAGTAAC
CACTATGCCAACCACATAGCCAAACAGCAGTCAGAAGAGGCATTTCTCAG
GAACAACAGAAGGCACCCCTGTTGTTGGGGGCTTCAATAGCAACGGGGGA
AGCAAGGTGTTAGGGCTCAAATATGAAGAGATTGACTGTCTCATAAACGAT
GAGCACACCATTAAGGGAGACGAGAGGGGAATGAAGTTTTCTTCCATTC
ACTTGGGTAGAGAAATACTTTGATGTTTATGGAAAAGTGGTTCGAGTATGA
CGGCTATGATCGATTTGAATTC::TCTCATAGCTATTCCAAAGTCTATGCACAGA
GAGCCCCTTATCACCTGATGGTGTGTTTATGTCCTTTGAAGGCTACAATGTGGA
AGTCCGAGACAGAGTCAAGTGCATAAGTGGGGTTGAAGGTGTACCTTTATCTACA
CAGTGGGGACCTCAAGGCTATTTCTACCCAATCCAGATTGCACAGTATGGGTAA
GTCACTACAGCAAGAATCTAACTGAAAAACCCCTCATATAGAGGTATATGAAA
CAGCAGAAGACAGGGACAAAAACAGCAAGCCCAATGACTGGACTGTGCCCAAGG
GCTGCTTTATGGCTAGTGTGGCTGATAAGTCAAGATTCACCAATGTTAAACAGTT
CATTGCTCCAGAAACCAGTGAAGGTGTATCCTTGCAACTGGGGAACACAAAAGA
TTTTATTATTTCAATTTGACCTCAAGTTCTTAACAAATGGAAGCGTGTCTGTGGTTC
TGGAGACGACAGAAAAGAATCAGCTCTTCACTGTACATTATGTCTCAAATACCCA

FIG. 1 CONT.

GCTAATTGCTTTTAAAGAAAGAGACATATACTATGGCATCGGGCCCAGAACATCA
TGGAGCACAGTTACCCGGGACCTGGTCACTGACCTCAGGAAAGGAGTGGGTCTTT
CCAACACAAAAGCTGTCAAGCCAACAAGAATAATGCCCAAGAAGGTGGTTAGGT
TGATTGCGAAAGGGAAGGGCTTCCTTGACAACATTACCATCTCTACCACAGCCCA
CATGGCTGCCTTCTTCGCTGCCAGTGACTGGCTGGTGAGGAACCAGGATGAGAAA
GGCGGCTGGCCGATTATGGTGACCCGTAAGTTAGGGGAAGGCTTCAAGTCTTTAG
AGCCAGGGTGGTACTCCGCCATGGCCCAAGGGCAAGCCATTTCTACATTAGTCAG
GGCCTATCTCTTAACAAAAGACCATATATTCCTCAATTCAGCTTTAAGGGCAACA
GCCCCTTACAAGTTTCTGTTCAGAGCAGCATGGAGTCAAGGCTGTGTTTATGAATA
AACATGACTGGTATGAAGAATATCCAACCTACACCTAGCTCTTTTGTTTTAAATGG
CTTTATGTATTCTTTAATTGGGCTGTATGACTTAAAAGAAACTGCAGGGGAAAAA
CTCGGGAAGAAGCGAGGTCCTTGTATGAGCGTGGCATGGAATCCCTTAAAGCC
ATGCTCCCCTTGTACGACACTGGCTCAGGAACCATCTATGACCTCCGGCACTTCA
TGCTTGGCATTGCCCCCAACCTGGCCCGCTGGGACTATCACACCACCCACATCAA
TCAACTGCAGCTGCTTAGCACCATTGATGAGTCCCCAATCTTCAAAGAATTTGTC
AAGAGGTGGAAGAGCTACCTTAAAGGCAGCCGGGCAAAGCACAACTAG

ATGCGTTGTTTGGCAGCTCGGGTCAACTATAAGACTTTGATTATCATCTGTGCGC
 TATTCACCTTTGGTCACAGTACTTTTGTGGAATAAGTGTTCCAGCGACAAAGCAAT
 CCAGTTTCCTCGGCACCTTGAGTAGTGGATTGAGAGTGGATTAGAAAAAAGA
 TCAGCAGCATCTGAAAGTAACCACTATGCCAACCACATAGCCAAACAGCAGTCA
 GAAGAGGCATTTCTCAGGAACAACAGAAGGCACCCCTGTTGTTGGGGGCTTCA
 ATAGCAACGGGGGAAGCAAGGTGTTAGGGCTCAAATATGAAGAGATTGACTGTC
 TCATAAACGATGAGCACACCATTAAAGGGAGACGAGAGGGGAATGAAGTTTTCC
 TTCCATTCACTTGGGTAGAGAAATACTTTGATGTTTATGGAAAAGTGGTCCAGTA
 TGACGGCTATGATCGATTTGAATTCTCTCATAGCTATTCCAAAGTCTATGCACAG
 AGATCACCTTATCACCCCTGACGGTGTGTTTATGTCCTTTGAAGGCTACAATGTGG
 AAGTCCGAGACAGAGTCAAATGTATAAGTGGAGTTGAAGGTGTGCCATTATCTAC
 CCAGTGGGGGCTCAAGGCTATTTCTACCCAATCCAGATTGCACAGTATGGGCTA
 AGTCATTACAGCAAGAATCTAACCGAGAAACCCCTCACATAGAAGTATATGAA
 ACAGCAGAAGACAGGGACAGAAACATCAGACCTAATGAATGGACTGTGCCCAAG
 GGGTGctCATGGCCAGTGTGGCAGACAAGTCTAGATCCACCAATGTTAAACAGTT
 TATTGCTCCAGAAACCAGTGAAGGTGTGTCTTTGCAGCTGGGAAACACAAAAGAC
 TTCATTATTTTCACTTGACCTCAAGCTTTTAACAAATGGGAGTGTGTCTGTGGTTCT
 GGAGACCACAGAAAAGAATCAGCTCTTCACTGTGCATTATGTCTCAAACACCCAG
 CTGATTGCTTTTCAAGAGACAGGGACATATACTACGGCATTGGGCCCAGAACTTCAT
 GGAGTACAGTTACCAGAGACCTGGTCACTGACCTCAGGAAAGGAGTGGGCCTTT
 CTAACACAAAAGCTGTCAAGCCAACCAAAATCATGCCCAAAAAGGTGGTTAGGT
 TGATTGCAAAAGGGAAGGGATTCTTGACAAACATTACCATCTCAACCACAGCCC
 ACATGGCTGCATTCTTTGCTGCAAGTGAAGTGGCTAGTGAGGAACCAGGATGAGAA
 AGGTGgctGGCCAATTATGGTGACCCGGAAGTTAGGGGAAGGGTTTAAATCTTTAG
 AACCAGGATGGTACTCTGCCATGGCACAAGGGCAAGCCATCTCTACCTTAGTCAG
 GGCCTATCTTCTAACGAAAGACTATGTATTCCTCAGTTCAGCTTTAAGGGCAACA
 GCCCCATACAAGTTTCCGTCAGAGCAGCATGGAGTTAAAGCCGTGTTTCATGAATA
 AACATGACTGGTATGAAGAATATCCAACCACACCTAGCTCTTTTGTTTTAAATGG
 CTTTATGTATTCTTTAATTGGGCTGTATGACCTAAAAGAAACAGCAGGGGAGACA

FIG. 2

CTTGGGAAAGAAGCAAGGTCCTTGTACGAGCGCGGCATGGAATCTCTTAAAGCC
ATGCTGCCCTTGTATGATACTGGCTCCGGGACCATCTATGACCTCCGCCACTTCA
TGCTTGGCATTGCTCCCAACCTGGCCCCGCTGGGACTATCACACCACCCACATTAA
CCAGCTGCAGCTGCTCAGCACCATCGATGAGTCCCCAATCTTCAAAGAATTTGTC
AAGAGGTGGAAAAGCTACCTTAAAGGCAGTAGGGCAAAGCACAAC

FIG. 2 CONT'D

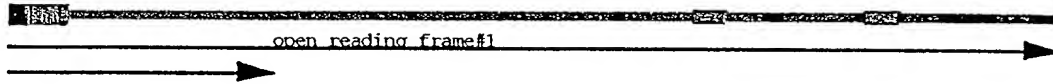
MetArgCysLeuAlaAlaArgValAsnTyrLysThrLeuIleIleIleCysAlaLeuPheThrLeuValThrValLe
 uLeuTrpAsnLysCysSerSerAspLysAlaIleGlnPheProArgHisLeuSerSerGlyPheArgValAspGlyL
 euGluLysArgSerAlaAlaSerGluSerAsnHisTyrAlaAsnHisIleAlaLysGlnGlnSerGluGluAlaPheP
 roGlnGluGlnGlnLysAlaProProValValGlyGlyPheAsnSerAsnGlyGlySerLysValLeuGlyLeuLy
 sTyrGluGluIleAspCysLeuIleAsnAspGluHisThrIleLysGlyArgArgGluGlyAsnGluValPheLeuP
 roPheThrTrpValGluLysTyrPheAspValTyrGlyLysValValGlnTyrAspGlyTyrAspArgPheGluP
 heSerHisSerTyrSerLysValTyrAlaGlnArgSerProTyrHisProAspGlyValPheMetSerPheGluGly
 TyrAsnValGluValArgAspArgValLysCysIleSerGlyValGluGlyValProLeuSerThrGlnTrpGlyPr
 oGlnGlyTyrPheTyrProIleGlnIleAlaGlnTyrGlyLeuSerHisTyrSerLysAsnLeuThrGluLysProPro
 HisIleGluValTyrGluThrAlaGluAspArgAspArgAsnIleArgProAsnGluTrpThrValProLysGlyC
 ysPheMetAlaSerValAlaAspLysSerArgSerThrAsnValLysGlnPheIleAlaProGluThrSerGluGly
 ValSerLeuGlnLeuGlyAsnThrLysAspPheIleIleSerPheAspLeuLysLeuLeuThrAsnGlySerValSe
 rValValLeuGluThrThrGluLysAsnGlnLeuPheThrValHisTyrValSerAsnThrGlnLeuIleAlaPhe
 ArgAspArgAspIleTyrTyrGlyIleGlyProArgThrSerTrpSerThrValThrArgAspLeuValThrAspLe
 uArgLysGlyValGlyLeuSerAsnThrLysAlaValLysProThrLysIleMetProLysLysValValArgLeuI
 eAlaLysGlyLysGlyPheLeuAspAsnIleThrIleSerThrThrAlaHisMetAlaAlaPhePheAlaAlaSerA
 spTrpLeuValArgAsnGlnAspGluLysGlyGlyTrpProIleMetValThrArgLysLeuGlyGluGlyPheLy
 sSerLeuGluProGlyTrpTyrSerAlaMetAlaGlnGlyGlnAlaIleSerThrLeuValArgAlaTyrLeuLeuT
 hrLysAspTyrValPheLeuSerSerAlaLeuArgAlaThrAlaProTyrLysPheProSerGluGlnHisGlyVal
 LysAlaValPheMetAsnLysHisAspTrpTyrGluGluTyrProThrThrProSerSerPheValLeuAsnGlyP
 heMetTyrSerLeuIleGlyLeuTyrAspLeuLysGluThrAlaGlyGluThrLeuGlyLysGluAlaArgSerLe
 uTyrGluArgGlyMetGluSerLeuLysAlaMetLeuProLeuTyrAspThrGlySerGlyThrIleTyrAspLeu
 ArgHisPheMetLeuGlyIleAlaProAsnLeuAlaArgTrpAspTyrHisThrThrHisIleAsnGlnLeuGlnLe
 uLeuSerThrIleAspGluSerProIlePheLysGluPheValLysArgTrpLysSerTyrLeuLysGlySerArgAl
 aLysHisAsn

FIG. 3

Fig. 4

sig seq-TM
conserved
peptide seq
hotspot

hydrophobic and
conserved peptide seq.
hotspots



Hypothetical orientation, if inserted into golgi

cytosol->lumen----->cytosol----->lumen

Key:

signal sequence,
highly hydrophobic
transmembrane (TM)
sequence

Hydrophobic
transmembrane (TM)
or buried sequence

most conserved peptide sequence
(>50% similarity to C elegans 71.9
KD hypothetical protein;
38% similarity to Methanococcus
hypothetical protein). Note:
peptide identity between mouse,
bovine and human > 95%!

Which in text appears:

mouse C5-e... 1 MRCLAARVNY KTLLITIDALF TLVTVLLWNK QSSDKAIQFP RHLSSGFRVD
51 GLEKRSAASE SNHYANHI AK QQSEEAFFQE QKAPPVVG FNSNGGSKVL
101 GLKYEEI DQL INDEHTI KGR REGNEVFLPF TWVEKYFDVY GKVVQYDGYD
151 RFEFSHSYSK VYAQRSPYHP DGVFMSFEGY NVEVRDRVKQ I SGVEGVPLS
201 TQWGPOGYFY PI QIAQYGLS HYSKNLTEKP PHIEVYETA DRDRNIRPNE
251 WTVPKG CFMA SVADKSRSTN VKQFIAPETS EGVSLQLGNT KDFIISFDLK
301 LLTNGSVSVV LETTEKNQLF TVHYVSNTQL IAFRDRDIYY GIGPRTSWST
351 VTRDLVTDLR KGVGLSNTKA VKPTKIMPKK VVRLIAKGKG FL DNITISTT
401 AHMAAFFAAS DWLVRNQDEK GGWPI MVTRK LGEGFKSLEP GWYSAMAQQQ
451 AISTLVRAYL LTKDYVFLSS ALRATAPYKF PSEQHGKAV FMNKHDWYEE
501 YP ITPSSFVL NGFMYS LI GL YDLKETAGET LGKEARSLYE RGMESLKAML
551 FLYDTSSGTI YDLRHFMLSI APNLARWBYH TTHINQLQLL STIDESPIFK
601 EFVKRWKSYL KGSRAKHN

FIG. 5

FIG. 6A

First active tagged recombinant (bovine) C5
(specific activity 5×10^5 cpm/mg/h)



The most active recombinant (full mouse) C5
(specific activity 2×10^5 cpm/mg/h)



Chimeric construct
(preliminary data indicate activity is 87% of full mouse):



Truncated mouse
(preliminary data indicate activity is same as first bovine construct):

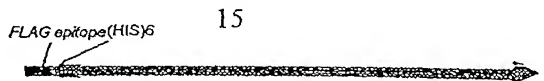
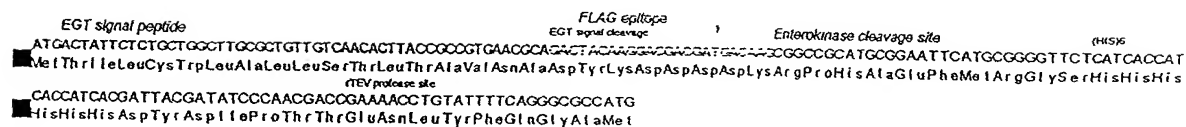


FIG. 6B



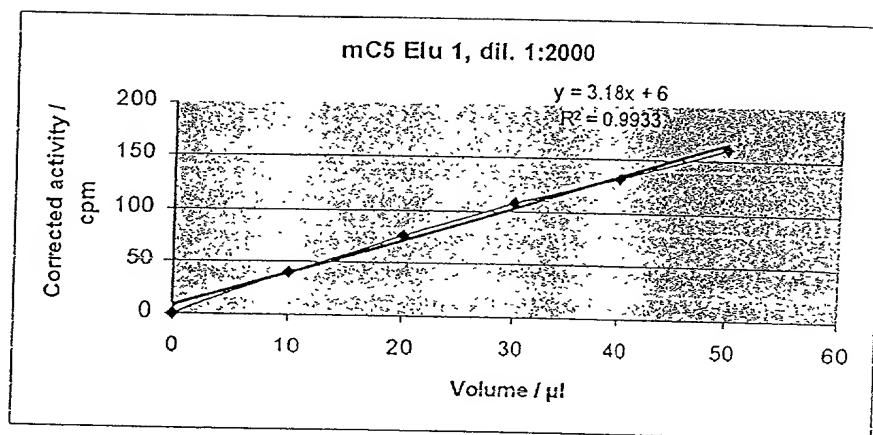


FIG. 7

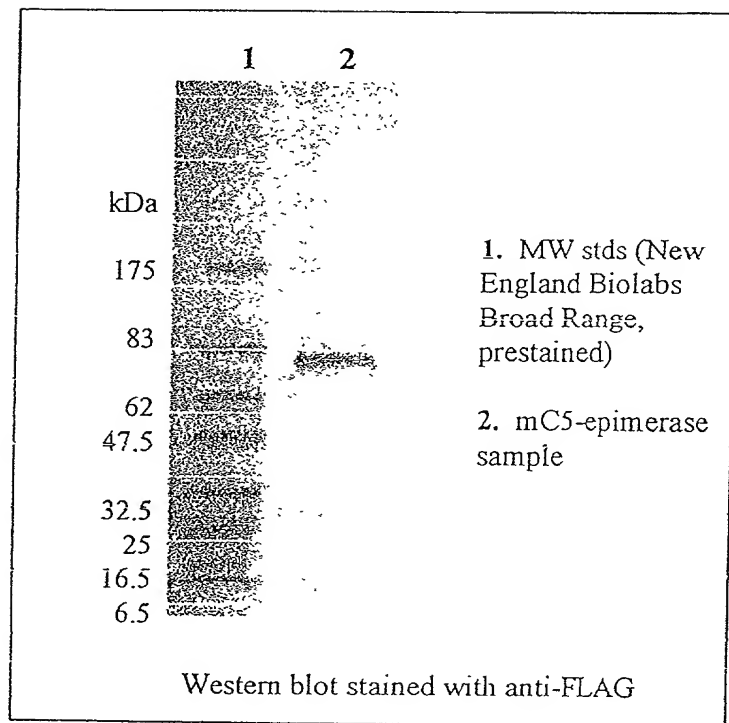


FIG. 8

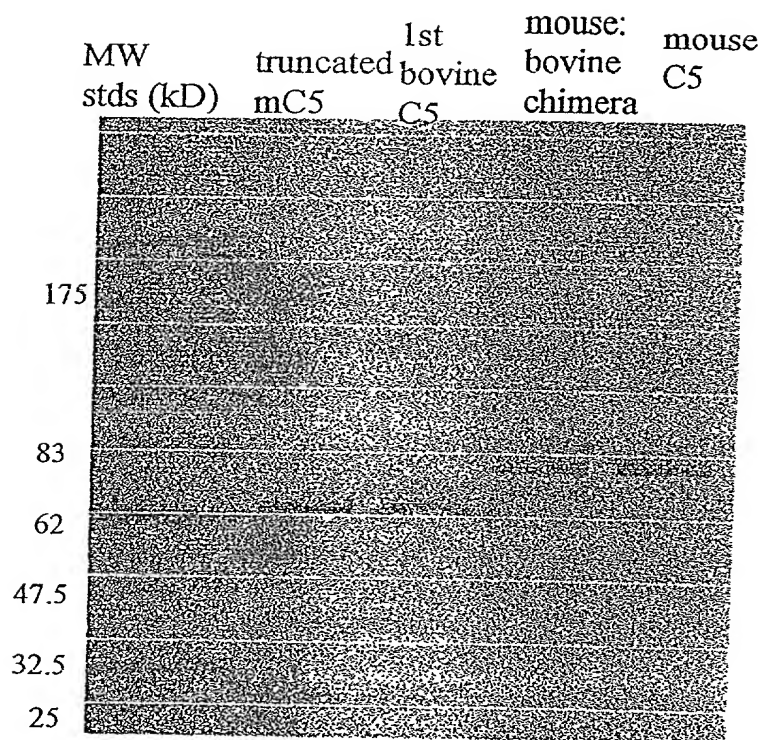


FIG. 9